	Type	н	# Hits	Search Text	DBs	Time Stamp	Comm	Error Defin ition	Er ro rs
1	BRS	L1	9590	epidermal adj growth adj factor	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/03/0 4 18:49			0
. 2	BRS	L2	1004	(epidermal adj growth adj factor) same modif\$7	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/03/0 4 18:36			0
3	BRS	ГЭ	297	laminin adj receptor	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/03/0 4 18:36			0
4	BRS	L4	139	(laminin adj receptor) same (antagonist or agonist or binding)	UB; PO; T	2003/03/0 4 18:37			0
5	BRS	LS	Н	2 same 4	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/03/0 4 18:37			0
9	BRS	L6	18	(epidermal adj growth adj factor) same (33-42)	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/03/0 4 18:59			0
7	BRS	L7	0	6 same modif\$7	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/03/0 4 18:58			0
æ	BRS	L8	0	6 same (tic or citrulline)	UB; PO; T	2003/03/0 4 18:59			0
0	BRS	L9	ĸ	(laminin adj receptor) same (antagonist or agonist) same retinopathy	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/03/0 4 19:02			0

	Type	# 1	Hit	Search Text	DBs	Time Stamp	Comm	Error Defin ition	ro rs
10	BRS	L10	0	(laminin adj receptor) same (antagonist or agonist) same (endothelial adj cell adj wounding)	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/03/0 4 19:02			0
11	BRS	L11	343	nelson adj john.in.	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/03/0 4 19:04			0
12	BRS	L12	59	walker adj brian.in.	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/03/0 4 19:04			0
13	BRS	L13	3	mcferran adj neil.in.	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/03/0 4 19:04			0
14	BRS	L14	2	harriott adj patrick.in.	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/03/0 4 19:05			0
15	BRS	L17	Н	(11 or 12 or 13 or 14) and 4	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/03/0 4 19:06			. 0

=> d his

## (FILE 'HOME' ENTERED AT 19:09:46 ON 04 MAR 2003)

FILE 'MEDLINE, CAPLUS, BIOSIS, EMBASE, SCISEARCH, AGRICOLA' ENTERED AT

19:10:12 ON 04 MAR 2003

- L1 136979 S (EPIDERMAL GROWTH FACTOR)
- L2 29 S L1 (P) (33-42)
- L3 4197 S L1 (P) MODIF?
- L4 1 S L2 (P) MODIF?
- L5 4070 S LAMININ RECEPTOR
- L6 1296 S L5 (P) (ANTAGONIST OR AGONIST OR BINDING)
- L7 9 S L6 (P) (L2 OR L3)
- L8 4 DUPLICATE REMOVE L7 (5 DUPLICATES REMOVED)
- L9 3 S L8 NOT L4
- L10 5 S L3 (P) (TIC OR CITRULLINE)
- L11 1 DUPLICATE REMOVE L10 (4 DUPLICATES REMOVED)
- L12 1 S L11 NOT L4
- L13 71189 S RETINOPATHY OR (ENDOTHELIAL CELL WOUNDING)
- L14 6 S L13 (P) L6
- L15 1 S L14 (P) L1
- L16 0 S L15 NOT L4

 $<sup>\</sup>Rightarrow$  log y

FILE 'HOME' ENTERED AT 19:09:46 ON 04 MAR 2003 => file medline caplus biosis embase scisearch agricola COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION 0.21 FULL ESTIMATED COST 0.21 FILE 'MEDLINE' ENTERED AT 19:10:12 ON 04 MAR 2003 FILE 'CAPLUS' ENTERED AT 19:10:12 ON 04 MAR 2003 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS) FILE 'BIOSIS' ENTERED AT 19:10:12 ON 04 MAR 2003 COPYRIGHT (C) 2003 BIOLOGICAL ABSTRACTS INC. (R) FILE 'EMBASE' ENTERED AT 19:10:12 ON 04 MAR 2003 COPYRIGHT (C) 2003 Elsevier Science B.V. All rights reserved. FILE 'SCISEARCH' ENTERED AT 19:10:12 ON 04 MAR 2003 COPYRIGHT (C) 2003 Institute for Scientific Information (ISI) (R) FILE 'AGRICOLA' ENTERED AT 19:10:12 ON 04 MAR 2003 => s (epidermal growth factor) 136979 (EPIDERMAL GROWTH FACTOR) => s 11 (p) (33-42)29 L1 (P) (33-42) => s l1 (p) modif? 4197 L1 (P) MODIF?

=> s 12 (p) modif?

1 L2 (P) MODIF?

=> d 14 1 ibib

ANSWER 1 OF 1 CAPLUS COPYRIGHT 2003 ACS ACCESSION NUMBER: 1999:691122 CAPLUS

DOCUMENT NUMBER:

131:295932

TITLE:

Peptide fragments of murine epidermal growth factor as laminin receptor targets for treatment of angiogenic

diseases

INVENTOR(S): Nelson, John; Walker, Brian; McFerran, Neil; Harriott,

Patrick

PATENT ASSIGNEE(S): The Queen's University of Belfast, UK SOURCE:

PCT Int. Appl., 35 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.				KIND DATE					APPLICATION NO.						DATE			
WO	9954	356		A1 19991028					WO 1999-GB1211						19990421			
	W:	ΑL,	AM,	AT,	AU,	ΑZ,	BA,	BB,	BG,	BR,	BY,	CA,	CH,	CN,	CU,	CZ,	DE,	
		DK,	EE,	ES,	FI,	GB,	GD,	GE,	GH,	GM,	HR,	HU,	ID,	ΙL,	IN,	IS,	JP,	
		KΕ,	KG,	ΚP,	KR,	ΚZ,	LC,	LK,	LR,	LS,	LT,	LU,	LV,	MD,	MG,	MK,	MN,	
		MW,	MX,	NO,	NZ,	PL,	PT,	RO,	RU,	SD,	SE,	SG,	SI,	SK,	SL,	ΤJ,	TM,	
		TR,	TT,	UA,	ŪĠ,	US,	UΖ,	VN,	YU,	ZA,	ZW,	AM,	ΑZ,	BY,	KG,	KZ,	MD,	
		RU,	ТJ,	TM														
•	RW:	GH,	GM,	KΕ,	LS,	MW,	SD,	SL,	SZ,	ŪĠ,	ZW,	AT,	BE,	CH,	CY,	DE,	DK,	
		ES,	FI,	FR,	GB,	GR,	ΙE,	IT,	LU,	MC,	NL,	PT,	SE,	BF,	ВJ,	CF,	CG,	
		CI,	CM,	GΑ,	GN,	GW,	ML,	MR,	ΝE,	SN,	TD,	TG				•		
AU 9936168			A1 19991108					AU 1999-36168						19990421				
EP 1073679			A1 20010207					EP 1999-918126					19990421					

```
AT, BE, CH, DE, DK, ES, R, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, FI
PRIORITY APPLN. INFO.:
                                        GB 1998-8407
                                                         Α
                                                            19980422
                                        WO 1999-GB1211
                                                         W
                                                            19990421
                               THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS
REFERENCE COUNT:
                         2
                               RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
=> d his
     (FILE 'HOME' ENTERED AT 19:09:46 ON 04 MAR 2003)
     FILE 'MEDLINE, CAPLUS, BIOSIS, EMBASE, SCISEARCH, AGRICOLA' ENTERED AT
     19:10:12 ON 04 MAR 2003
         136979 S (EPIDERMAL GROWTH FACTOR)
L1
L_2
             29 S L1 (P) (33-42)
           4197 S L1 (P) MODIF?
1.3
              1 S L2 (P) MODIF?
=> s laminin receptor
          4070 LAMININ RECEPTOR
=> s l5 (p) (antagonist or agonist or binding)
          1296 L5 (P) (ANTAGONIST OR AGONIST OR BINDING)
=>'s 16 (p) (12 or 13)
             9 L6 (P) (L2 OR L3)
=> duplicate remove 17
DUPLICATE PREFERENCE IS 'MEDLINE, CAPLUS, BIOSIS, EMBASE, SCISEARCH'
KEEP DUPLICATES FROM MORE THAN ONE FILE? Y/(N):n
PROCESSING COMPLETED FOR L7
              4 DUPLICATE REMOVE L7 (5 DUPLICATES REMOVED)
=> s 18 not 14
             3 L8 NOT L4
=> d 19 1-3 ibib abs
     ANSWER 1 OF 3
                       MEDLINE
ACCESSION NUMBER:
                    96421617
                                 MEDLINE
DOCUMENT NUMBER:
                               PubMed ID: 8824265
                    96421617
TITLE:
                    Murine epidermal growth factor peptide (33-42) binds to a
                    YIGSR-specific laminin receptor on both tumor and
                    endothelial cells.
                    Nelson J; Scott W N; Allen W E; Wilson D J; Harriott P;
AUTHOR:
                    McFerran N V; Walker B
CORPORATE SOURCE:
                    Centre for Peptide and Protein Engineering, School of
                    Biology and Biochemistry, The Queen's University of
                    Belfast, Belfast BT9 7BL, Northern Ireland, United Kingdom.
SOURCE:
                    JOURNAL OF BIOLOGICAL CHEMISTRY, (1996 Oct 18) 271 (42)
                    26179-86.
                    Journal code: 2985121R. ISSN: 0021-9258.
PUB. COUNTRY:
                    United States
DOCUMENT TYPE:
                    Journal; Article; (JOURNAL ARTICLE)
LANGUAGE:
                    English
FILE SEGMENT:
                    Priority Journals
ENTRY MONTH:
                    199611
ENTRY DATE:
                    Entered STN: 19961219
                    Last Updated on STN: 20000303
                    Entered Medline: 19961126
    A laminin- ***antagonist*** peptide, comprising amino acids ***33***
       ***42***
                              ***epidermal*** ***growth***
                 of murine
                      (mEGF-( ***33*** - ***42*** )), interacts with a breast
       ***factor***
     cancer- and endothelial cell-associated receptor, which is specific for
     the laminin B1 chain sequence, CDPGYIGSR-NH2 (Lam.B1-(925-933)), and is
     immunologically similar to a previously described 67-kDa ***laminin***
       ***receptor*** . In whole cell receptor assays, mEGF-( ***33***
       ***42*** ), Lam. B1-(925-933), and laminin all have IC50 values for
     displacement of 125I-laminin in the range 1-5 nM. Cell attachment to
     solid-phase laminin is also blocked by all three ligands, but in contrast
     to the receptor assays, mEGF-( ***33*** - ***42*** ) or
```

Lam.B1-(925-933), while equipotert with each other, were less effective than laminin. The concentration of the peptides required to problem half-maximal inhibition of attachment were in the range 230-390 nm, but those for laminin were 1000-fold lower, in the range 0.2-0.3 nm. Like laminin, solid-phase mEGF-( \*\*\*33\*\*\* - \*\*\*42\*\*\* ) supports cell attachment, and this ability is blocked by anti-67-kDa receptor antibodies. Modeling studies suggest that both peptides present a tyrosyl and an arginyl residue on the same face of a right-handed helical fold with elliptical cross-section.

L9 ANSWER 2 OF 3 MEDLINE

ACCESSION NUMBER: 86199471 MEDLINE

DOCUMENT NUMBER: 86199471 PubMed ID: 3457945

TITLE: Chemotaxis of human gingival epithelial cells to laminin. A

mechanism for epithelial cell apical migration.

AUTHOR: Terranova V P; Lyall R M

SOURCE: JOURNAL OF PERIODONTOLOGY, (1986 May) 57 (5) 311-7.

Journal code: 8000345. ISSN: 0022-3492.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Dental Journals; Priority Journals

ENTRY MONTH: 198606

ENTRY DATE: Entered STN: 19900321

Last Updated on STN: 19900321 Entered Medline: 19860613

AB Laminin, a large glycoprotein (Mr = 10(6)) and a major component of basement membrane, is shown here to be a potent chemoattractant for human gingival epithelial cells. Laminin stimulated chemotaxis and chemokinesis of gingival epithelial cells in the \*\*\*modified\*\*\* Boyden chamber \*\*\*laminin\*\*\* assay. This effect appeared to be \*\*\*receptor\*\*\* mediated. Gingival epithelial cells were shown to bind laminin (Kd = 2.0 \*\*\*binding\*\*\* sites per cell. Antilaminin nM) with 10,000 to 30,000 \*\*\*binding\*\*\* , inhibited the antibody, which inhibited laminin chemotactic response of epithelial cells to laminin, while antifibronectin was without effect. Fibronectin was not as potent a chemoattractant as laminin. Other biological response \*\*\*modifiers\*\*\* were also tested;

\*\*\*factor\*\*\* were active as chemoattractants, although not as effective in inducing chemotaxis as laminin. The data indicate that laminin and other components of basement membrane may be important in regulating the migration and growth of gingival epithelial cells.

\*\*\*epidermal\*\*\*

\*\*\*growth\*\*\*

L9 ANSWER 3 OF 3 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.

ACCESSION NUMBER: 2002:111703 BIOSIS DOCUMENT NUMBER: PREV200200111703

of these, Type IV collagen and

TITLE: Synthetic peptides interacting with the 67-kd laminin

receptor can reduce retinal ischemia and inhibit

hypoxia-induced retinal neovascularization.

AUTHOR(S): Gebarowska, Dorota; Stitt, Alan W. (1); Gardiner, Thomas

A.; Harriott, Patrick; Greer, Brett; Nelson, John

(1) Center of Ophthalmology and Vision Science, The Queen's

University of Belfast, Royal Victoria Hospital, Belfast,

BT12 6BA: a.stitt@qub.ac.uk UK

SOURCE: American Journal of Pathology, (January, 2002) Vol. 160,

No. 1, pp. 307-313. http://ajp.amjpathol.org/. print.

ISSN: 0002-9440.

DOCUMENT TYPE: Article LANGUAGE: English

CORPORATE SOURCE:

The high-affinity 67-kd laminin receptor (67LR) is expressed by proliferating endothelial cells during retinal neovascularization. The role of 67LR has been further examined experimentally by administration of selective 67LR agonists and antagonists in a murine model of proliferative retinopathy. These synthetic 67LR ligands have been previously shown to stimulate or inhibit endothelial cell motility in vitro without any direct effect on proliferation. In the present study, a fluorescently labeled 67LR antagonist (EGF33-42) was injected intraperitoneally into mice and its distribution in the retina was assessed by confocal scanning laser microscopy. Within 2 hours this peptide was localized to the retinal vasculature, including preretinal neovascular complexes, and a significant amount had crossed the blood retinal barrier. For up to 24 hours postinjection, the peptide was still present in the retinal vascular walls

and, to a lesser extent, in the pural retina. Non-labeled EGF33-42 significantly inhibited pre-retion neovascularization in compared to controls treated with phosphate-buffered saline or scrambled peptrde (P < 0.0001). The agonist peptide (Lambeta1925-933) also significantly inhibited proliferative retinopathy; however, it caused a concomitant reduction in retinal ischemia in this model by promoting significant revascularization of the central retina (P < 0.001). Thus, 67LR appears to be an important target receptor for the modulation of retinal neovascularization. Agonism of this receptor may be valuable in reducing the hypoxia-stimulated release of angiogenic growth factors which drives retinal angiogenesis.

## => d his (FILE 'HOME' ENTERED AT 19:09:46 ON 04 MAR 2003) FILE 'MEDLINE, CAPLUS, BIOSIS, EMBASE, SCISEARCH, AGRICOLA' ENTERED AT 19:10:12 ON 04 MAR 2003 L1136979 S (EPIDERMAL GROWTH FACTOR) 29 S L1 (P) (33-42) L24197 S L1 (P) MODIF? L3 1 S L2 (P) MODIF? L4L5 4070 S LAMININ RECEPTOR 1296 S L5 (P) (ANTAGONIST OR AGONIST OR BINDING) L6 9 S L6 (P) (L2 OR L3) L7 L8 4 DUPLICATE REMOVE L7 (5 DUPLICATES REMOVED) L9 3 S L8 NOT L4 => s 13 (p) (tic or citrulline) 5 L3 (P) (TIC OR CITRULLINE) => duplicate remove 110 DUPLICATE PREFERENCE IS 'MEDLINE, CAPLUS, BIOSIS, EMBASE, SCISEARCH' KEEP DUPLICATES FROM MORE THAN ONE FILE? Y/(N):n PROCESSING COMPLETED FOR L10 1 DUPLICATE REMOVE L10 (4 DUPLICATES REMOVED) => s l11 not l4 1 L11 NOT L4 L12=> d l12 1 ibib abs L12 ANSWER 1 OF 1 MEDLINE ACCESSION NUMBER: 93248272 MEDLINE DOCUMENT NUMBER: 93248272 PubMed ID: 7683432 TITLE: Differential regulation of inducible nitric oxide synthase by fibroblast growth factors and transforming growth factor beta in bovine retinal pigmented epithelial cells: inverse correlation with cellular proliferation. AUTHOR: Goureau O; Lepoivre M; Becquet F; Courtois Y CORPORATE SOURCE: Unite de Recherches Gerontologiques, Institut National de la Sante et de la Recherche Medicale U118, Paris, France. SOURCE: PROCEEDINGS OF THE NATIONAL ACADEMY OF SCIENCES OF THE UNITED STATES OF AMERICA, (1993 May 1) 90 (9) 4276-80. Journal code: 7505876. ISSN: 0027-8424. PUB. COUNTRY: United States DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE) LANGUAGE: English FILE SEGMENT: Priority Journals ENTRY MONTH: 199306 ENTRY DATE: Entered STN: 19930618 Last Updated on STN: 20000303 Entered Medline: 19930601 AB Bovine retinal pigmented epithelial (RPE) cells express, after activation

with interferon gamma (IFN-gamma) and lipopolysaccharide (LPS), an inducible nitric oxide synthase (NOS). Experiments were done to investigate the effects of the transforming growth factor beta 1,

\*\*\*factor\*\*\*

\*\*\*growth\*\*\*

\*\*\*epidermal\*\*\*

factors (FGFs), which are abundant in the retina, on NOS activity. Transforming growth factor beta 1 slightly increases the production of nitrite, an oxidation product of NO, induced by LPS plus IFN-gamma,

, and fibroblast growth

```
whereas acidic and basic FGFs markedly inhibit the nitrite release due to LPS/IFN-gamma in a concentration ependent manner, and ***epid al***

***growth*** ***factor*** did not ***modify***
     LPS/IFN-gamma-induced NOS activity. The growth factors alone did not
     stimulate nitrite release. We have attempted to elucidate the mechanism of
     FGF inhibition. Results with heparin, suramin, and tyrphostin suggest
     involvement of the high-affinity receptor for FGF in its inhibition of
     LPS/IFN-gamma-stimulated NOS activity. Continued stimulation of RPE cells
     with LPS/IFN-gamma was essential for the induction of NO synthesis, and
     maximal inhibition was obtained when FGF was present during stimulation
     with LPS/IFN-gamma, suggesting that FGF inhibits NOS induction.
     Furthermore, an antiproliferative action of NO was demonstrated by an
     inverse correlation between the amounts of nitrite or
     produced in response to different stimuli (LPS/IFN-gamma or LPS/IFN-gamma
     with growth factors) and the level of cellular proliferation. Similar
     inhibition of growth was obtained when RPE cells were incubated with an NO
     donor, sydnonimide. Because NO acts as a cytotoxic compound in the retina,
     FGF, by inhibiting the induction of NOS in RPE cells, may have beneficial
     effects in protecting the retina from cytokine and endotoxin-mediated
     tissue damage.
=> d his
     (FILE 'HOME' ENTERED AT 19:09:46 ON 04 MAR 2003)
     FILE 'MEDLINE, CAPLUS, BIOSIS, EMBASE, SCISEARCH, AGRICOLA' ENTERED AT
     19:10:12 ON 04 MAR 2003
         136979 S (EPIDERMAL GROWTH FACTOR)
              29 S L1 (P) (33-42)
           4197 S L1 (P) MODIF?
               1 S L2 (P) MODIF?
           4070 S LAMININ RECEPTOR
           1296 S L5 (P) (ANTAGONIST OR AGONIST OR BINDING)
               9 S L6 (P) (L2 OR L3)
               4 DUPLICATE REMOVE L7 (5 DUPLICATES REMOVED)
               3 S L8 NOT L4
               5 S L3 (P) (TIC OR CITRULLINE)
               1 DUPLICATE REMOVE L10 (4 DUPLICATES REMOVED)
               1 S L11 NOT L4
=> s retinopathy or (endothelial cell wounding)
   4 FILES SEARCHED...
         71189 RETINOPATHY OR (ENDOTHELIAL CELL WOUNDING)
=> s 113 (p) 16
             6 L13 (P) L6
=> s 114 (p) 11
             1 L14 (P) L1
=> s 115 not 14
             0 L15 NOT L4
=> d his
     (FILE 'HOME' ENTERED AT 19:09:46 ON 04 MAR 2003)
     FILE 'MEDLINE, CAPLUS, BIOSIS, EMBASE, SCISEARCH, AGRICOLA' ENTERED AT
     19:10:12 ON 04 MAR 2003
         136979 S (EPIDERMAL GROWTH FACTOR)
             29 S L1 (P) (33-42)
           4197 S L1 (P) MODIF?
              1 S L2 (P) MODIF?
           4070 S LAMININ RECEPTOR
           1296 S L5 (P) (ANTAGONIST OR AGONIST OR BINDING)
              9 S L6 (P) (L2 OR L3)
              4 DUPLICATE REMOVE L7 (5 DUPLICATES REMOVED)
              3 S L8 NOT L4
              5 S L3 (P) (TIC OR CITRULLINE)
              1 DUPLICATE REMOVE L10 (4 DUPLICATES REMOVED)
              1 S L11 NOT L4
```

L1

L2

L3

L5

L6 L7

L8

L9

L10 1.11

L13

L14

L15

1.16

L1

L2

L3

L4 $L_5$ 

L<sub>6</sub>

L7

L8

1.9

L10

L11

L12

L15 1 S L14 (P) L1
L16 0 S L15 NOT L4

=> log y
COST IN U.S. DOLLARS
SINCE FILE TOTAL
ENTRY SESSION
FULL ESTIMATED COST 46.14 46.35

71189 S RETINOPATHY OR (ENDOTHELIAL CELL WOUNDING)

STN INTERNATIONAL LOGOFF AT 19:18:07 ON 04 MAR 2003

6 S L13 (P) L6

L13

L14